



Docket No.: 7853-234-999  
Serial No.: 09/829,495  
Inventor(s): BUSFIELD ET AL.  
Title: "GLYCOPROTEIN VI AND USES THEREOF"

09829495.080102

#J

	M	S	P	S	P	T	A	L	F	C	L	11
GGAGTCGACCCACGCGTCCGCGAGGCTGAGGAACC	ATG	TCT	CCA	TCC	CCG	ACC	GCC	CTC	TTC	TGT	CTT	68
G L C L G R V P A Q S G P L P K P S L Q	31											
GGG CTG TGT CTG GGG CGT GTG CCA GCG CAG AGT GGA CCG CTC CCC AAG CCC TCC CTC CAG	128											
A L P S S L V P L E K P V T L R C Q G P	51											
GCT CTG CCC AGC TCC CTG GTG CCC CTG GAG AAG CCA GTG ACC CTC CGG TGC CAG GGA CCT	188											
P G V D L Y R L E K L S S S R Y Q D Q A	71											
CCG GGC GTG GAC CTG TAC CGC CTG GAG AAG CTG AGT TCC AGC AGG TAC CAG GAT CAG GCA	248											
V L F I P A M K R S L A G R Y R C S Y Q	91											
GTC CTC TTC ATC CCG GCC ATG AAG AGA AGT CTG GCT GGA CGC TAC CGC TGC TCC TAC CAG	308											
N G S L W S L P S D Q L E L V A T G V F	111											
AAC GGA AGC CTC TGG TCC CTG CCC AGC GAC CAG CTG GAG CTC GTT GCC ACG GGA GTT TTT	368											
A K P S L S A Q P G P A V S S G G D V T	131											
GCC AAA CCC TCG CTC TCA GCC CAG CCC GGC CCG GCG GTG TCG TCA GGA GGG GAC GTA ACC	428											
L Q C Q T R Y G F D Q F A L Y K E G D P	151											
CTA CAG TGT CAG ACT CGG TAT GGC TTT GAC CAA TTT GCT CTG TAC AAG GAA GGG GAC CCT	488											
A P Y K N P E R W Y R A S F P I I T V T	171											
GCG CCC TAC AAG AAT CCC GAG AGA TGG TAC CGG GCT AGT TTC CCC ATC ATC ACG GTG ACC	548											
A A H S G T Y R C Y S F S S R D P Y L W	191											
GCC GCC CAC AGC GGA ACC TAC CGA TGC TAC AGC TTC TCC AGC AGG GAC CCA TAC CTG TGG	608											
S A P S D P L E L V V T G T S V T P S R	211											
TCG GCC CCC AGC GAC CCC CTG GAG CTT GTG GTC ACA GGA ACC TCT GTG ACC CCC AGC CGG	668											
L P T E P P S S V A E F S E A T A E L T	231											
TTA CCA ACA GAA CCA CCT TCC TCG GTA GCA GAA TTC TCA GAA GCC ACC GCT GAA CTG ACC	728											
V S F T N K V F T T E T S R S I T T S P	251											
GTC TCA TTC ACA AAC AAA GTC TTC ACA ACT GAG ACT TCT AGG AGT ATC ACC ACC AGT CCA	788											
K E S D S P A G P A R Q Y Y T K G N L V	271											
AAG GAG TCA GAC TCT CCA GCT GGT CCT GCC CGC CAG TAC TAC ACC AAG GGC AAC CTG GTC	848											
R I C L G A V I L I I L A G F L A E D W	291											
CGG ATA TGC CTC GGG GCT GTG ATC CTA ATA ATC CTG GCG GGG TTT CTG GCA GAG GAC TGG	908											
H S R R K R L R H R G R A V Q R P L P P	311											
CAC AGC CGG AGG AAG CGC CTG CGG CAC AGG GGC AGG GCT GTG CAG AGG CCG CTT CCG CCC	968											

FIG.1A



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L P P L P Q T R K S H G G Q D G G R Q D 331  
CTG CCG CCC CTC CCG CAG ACC CGG AAA TCA CAC GGG GGT CAG GAT GGA GGC CGA CAG GAT 1028

V H S R G L C S \* 340  
GTT CAC AGC CGC GGG TTA TGT TCA TGA 1055

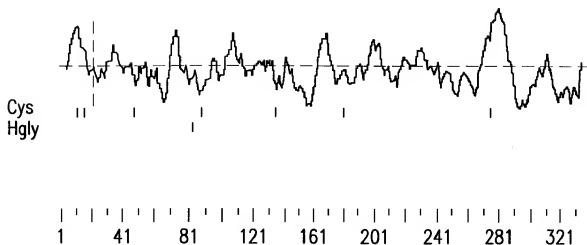
CCGCTGAACCCACGGCAGCGTCTGATCCAAAGGGAGGGATCATGGCATGGGAGGCAGCTCAAAGACTGGCGTGTGTGGAG 1134  
CGTGGAAGCAGGAGGGCAGAGGCTACAGCTGTGAAACGAGGCCATGCTGCCTCCTCCTGGTGTTCATCAGGGAGCCG 1213  
TTCGGCCAGTGTCTGTCTGTCTGTCTGCCTCTCTGTCTGAGGGCACCCCTCCATTGGGATGGAAGGAATCTGTGGAGAC 1292  
CCCATCCTCCTCCCTGCACACTGTGGATGACATGGTACCCTGGCTGGACCACATACTGGCCTCTTTCTTCAACCTCTCT 1371  
AATATGGGCTCCAGACGGATCTCTAAGGTTCCAGCTCTCAGGGTTGACTCTGTTCCATCCTCTGTGCAAAATCCTCCT 1450  
GTGCTTCCCTTTGGCCCTCTGTGCTCTTGTCTGGTTTTCCCAGAACTCTCACCCCTCACTCCATCTCCCACTGCGGTC 1529  
TAACAAATCTCCTTTCTGCTCTCTCAGAACGGGTCTTGCAGGCAGTTTGGGTATGTCAATTCATTTTCCTTAGTGTAAACT 1608  
AGCAGCTTGCCCGCTTCCCTTCACTTAGAAAAACAAGATCAGCCTGTGCAACATGGTGAACCTCATCTCTACCAACAA 1687  
AACAAAAAACACAAAAATTAGCCAGGTGTGGTGGTGCATCCCTATACTCCAGCAACTCGGGGGGCTGAGGTGGGAGA 1766  
ATGGCTTGAGCCTGGGAGGCAGAGGTTGCAGTGAGCTGAGATCACACCACTGCACCTAGCTCGGGTGACGAGCCCTGA 1845  
CCTGTGTCAAAAAATACAGGGATGAATATGTCAATTACCTGATTTGATCATAGACGTTGTATACATGTACTGCAAT 1924  
ATTGCTGTCCACCCATAAATATGTACAATTATGTATACATTTTTAAATCATAAAAATAAGATAATGAAAAAAAAA 2003  
AAAAAAAAAAAAAGGGCGGCGCTAGACTAGTCTAGAGAACA 2047

FIG.1B



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MSPSP TALFCLGLCLGRVPAQSGPLPKPSLQALPSSLVPLEKPVTLRCQGP PGVDLYRLE  
KLSSRYQDQAVLFIPAMKRSLAGRYRCSYQNGSLWSLPSDQLELVATGVFAKPSLSAQ  
GPAVSSGGDVTLCQQTRYGFDQFALYKEGDPAPYKNPERWYRASFPITVTAAHSGTYRC  
YSFSSRDPLYWSAPSDPLELVVTGTSVTPSRLPTEPPSSVAEFSEATAELTVSF TNKVF  
TETSRSIITSPKESDSPAGPARQYYTKGNLVRICLGAVILIIILAGFLAEDWHSRRKRLRH  
RGRAVQRPLPPLPPLPQTRKSHGGQDGRQDVHSRGLCS

FIG.2





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```

560      570      580      590      600      610      620
inputs  GAACCCAGCCACAGGTGGAGGTTACATGCTATTACTATTATATGAACACCCCCAGGTGTGGTCCCAC
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      GAATCCCGA-----GAGATGGTAC-CGGGCTAGT---TT-----CCCCAT-----CAT
      470              480      490              500

630      640      650      660      670      680      690
inputs  CCCAGTGACCCCTGGAGATTCTGCCCTCAGGCGTGTCTAGGAAGCCCTCCCTCCTGACCCCTGCAGGGCC
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      CACGGTGACCGCC-----GCCACAG-----
      510              520

700      710      720      730      740      750      760
inputs  CTGTCTGGCCCTGGGCAGAGCCTGACCTCCAGTGTGGCTCTGATGTGGCTACGACAGATTTGTTCT
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      -----CGGAACCTA---CCGATG-----CTACAGC-----TTCT
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      530              540              550

770      780      790      800      810      820      830
inputs  GTATAAGGAGGGGGAACGTGACTTCTCCAGCGCCCTGGCCAGCAGCCCAAGGCTGGGCTCTCCAGGGCC
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      -----CCAGCAG-----

840      850      860      870      880      890      900
inputs  AACTTCACCCCTGGGCCCTGTGAGCCCTCCACGGGGCCAGTACAGGTGCTATGGTGACACAACTCT
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      -----GGACCCA-----TACCT--
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      560

910      920      930      940      950      960      970
inputs  CCTCCGAGTGGTCGGCCCCAGCGACCCCTGAACATCCTGATGGCAGGACGATCTATGACACCGTCTC
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      -----GTGGTCGGCCCCAGCGACCCCTGGA-----GCT-----TGTG-----
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      570      580      590              600

980      990      1000      1010      1020      1030      1040
inputs  CCTGTGACGACAGCGGGCCACAGTGGCTCAGGAGAGAAGCTGACCTGCTGTGTCAGTCATGGTGG
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      ---GTCA-----CAGGAACCTCTGTGACC-----CCAGC-----CGGT-----
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      610      620              630

1050      1060      1070      1080      1090      1100      1110
inputs  CAGTTTGACACTTTCTTCTGACCAAGAAGGGGACGCCATCCCCACCTGCGTCTGAGATCAATGTACG
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      -----TACCAACAGAAC-----CA--CCTTCC-----TCG
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      640              650

1120      1130      1140      1150      1160      1170      1180
inputs  GAGCTCATAGATTACCAAGGCTGAATTCCTCATGAGTCTGTGACCTCAGCCACGCGGGGACCTACAGGTG
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      GTA-----GCAGAAITTC-----AGAGCCAC-----CGTGTA-----ACTG--A
      : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : : :
      660              670              680              690
  
```

FIG.3B



```

1190      1200      1210      1220      1230      1240      1250
inputs  CTACGGCTCATACAGCTCCAACCCCCACCTGCTGCTCTTCCCCAGTGAGCCCTGGAACTCATGGTCTCA
        C--CGTCTCATTCA--CAAAC-----AAAGTCTT--CACAA-----CTGAGACT--TCT--
                700                710                720                730

1260      1270      1280      1290      1300      1310      1320
inputs  GGACACTCTGGAGGCTCCAGCCTCCACCCACAGGGCCGCCCTCCACACCTGCTGGGAAGATACCTGG
        -----AGGAGTATC--ACCACCAGTCCAAAGGA--GTCAGACTCTCCAG--CTGG-----
                740                750                760                770

1330      1340      1350      1360      1370      1380      1390
inputs  AGSTTTTGATTGGGTCCTGGTGGCCTTCGCTGCTGCTCTTCTCTCTCTCTCTCTCTCTCTCCGACG
        -----TCCTGC-----CCGCCAGTA--CTACACCAAGG
                780                790                800

1400      1410      1420      1430      1440      1450      1460
inputs  TCAGCGCTCAGCAAAACACAGGACATCTGACCAGAGAAAAGACTGATTTTCAGCGTCTGCAGGGGCTGCG
        GCAAC-----CTGGTC-----CGGATAT--GCCTC-----GGGGCTG--
                810                820                830

1470      1480      1490      1500      1510      1520      1530
inputs  GAGACAGAGCCCAAGGACAGGGGCTGCTGAGGAGGTCCAGCCCAGCTGCTGACGCTCCAGGAAGAAAACC
        ----TGATCCTAATAA-----TCCTG--CGGGGGTTTCTG-----GCAGA--GGACTGG-----C
                840                850                860                870

1540      1550      1560      1570      1580      1590      1600
inputs  TCTATGCTGCCGTGAAGGACACACAGTCTGAGG--ACAGGGTGGAGCTGGACAGT--CAGAGCCACACAGAT
        AC-----AGCCG--GAGGAAGCG--CTGCGGCACAGGG--GCAGGGCTGTGCAGAGGCCGCT--
                880                890                900                910                920

1610      1620      1630      1640      1650      1660      1670
inputs  GAAGACCCCCAGGCAGTGACGTATGCCCCGGTGAAACTCCAGTCTAGGAGAGAAATGGCTCTCTCTC
        ---TCC-----GCCCTG-----CCGC-----C
                930                940

1680      1690      1700      1710      1720      1730      1740
inputs  CCTCTCACTGTCTGGGAATTCTTGACACAAAAGCAGACAGGTGGAAGAGGACAGGCAGATGGACAC
        CCTCC--CGCAGAC-----CCGGAATCA--CA--CGGG-----GGTCAGG--ATGGA--
                950                960                970                980

1750      1760      1770      1780      1790      1800      1810
inputs  TGAGGCTGCTGCATCTGAAGCCTCCAGGATGTGACCTACGCCAGCTGCACAGCTTGACCTTAGACGG
        ---GCC-----CGAC-----AGGATGTT-----CACAG-----CG-
                990                1000

1820      1830      1840      1850      1860      1870      1880
inputs  AAGGCAACTGAGCTCTCCATCCAGGAAGGGGAACCTCCAGCTGAGCCAGCATCTACGCCACTCTGG
        -----CGGGTTATG-----TTCA-----
                1010

1890
inputs  CCATCCAC
        -----
  
```

FIG.3C

**inputs**

MSPSP<sup>10</sup>TALFCLGLCLG-RVPAQSGPLPKPSLQALPSSLVPLEKPVTLRCQQPPGV<sup>20</sup>DLYRLEKLSSS-----  
:::  
MTPALTALLCLGLSLGPRTRVQAGFPFKPTLWAEPGSVISWGSGPVTIWCQGSLEAQYRLDKEGSEPLD<sup>30</sup>  
:::

**inputs**

R<sup>70</sup>YQ----DQAVLFIPAMKRSLAGRYCSYQNGSLWSLPSDQLVLATGVFAKPSLSAQPGAPVASGGDV<sup>80</sup>  
:::  
RNNPLEPKNKARFSIPSMTTEHHAGRYRCHYYSSAGWSEPSDPLELVMTGFYNKPTLSALPSPVVASGGNM<sup>90</sup>  
:::

**inputs**

TLQCQT-----RY-----  
:::  
TLRCGSQKGYYHFLVMKEGEHQLPRTLDSQQLHSGGFQALFPVGVPNPSHRWRFTCYYYYMMNTQPQW<sup>150</sup>SH<sup>160</sup>P  
:::

**inputs**

-----140150-----  
:::  
SDPLEILPSGVSRKPSLLTLQGVPVLAQGSQLTLCQCGSDVGVDYDRFVLYKEGERDFLRQPGQQPAGLSQAN<sup>180</sup>  
:::

**inputs**

-----APYK-----NP-----  
:::  
FTLGVPVPSHGQYRCYGAHNLSSWEAPSDDLNI<sup>220</sup>LMAQIYDTVSLSAQPGPTVASGENVTLLCQSWMQ<sup>230</sup>  
:::

**inputs**

-----170180190200-----  
:::  
FD<sup>360</sup>TLLTKEGAHPPLRLRSMYGAHKYQAEEFMSPVTSAHAGTYRCYGSYSNPHLLSFPSEPELMVSG<sup>370</sup>  
:::

**inputs**

<sup>210</sup>TSVTPSRLPTEPPSS--VAEFSEATAELTVSF<sup>220</sup>NKV-----TTETSRSTITSPKESD--SPAGA-<sup>240</sup>  
:::  
<sup>250</sup>HSGGSSLPPTGPPSPGLGRYLEVLIGVSVAFLVLL<sup>260</sup>LLLLRRQRHSKHRTSDORKTD<sup>270</sup>FQARGAAE<sup>280</sup>  
:::

**inputs**

<sup>270</sup>RQYYTKGNLVRICLAGVIL-----IILAGFLAEDW-----HSRRKR-----  
:::  
<sup>280</sup>TEPKDRGLLRRSSPAADVQEENLYAAVKDTQSEDRVELDSQSPHDEDPQAVTYAPVKHSSPRREMASPPS<sup>290</sup>  
:::

**inputs**

-----300310320330-----  
:::  
SLSGEFLTOKDRQVEEDROMDTEAAEAASEAQDV<sup>340</sup>VTYAQLHSLTLRRKATEPPPSQEGEPPEAFPSIYATLAI<sup>350</sup>  
:::

**inputs S**

## H

FIG.4



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      *->GesvtLtCsvgsgfgppgsvtWyfkngk.lgpsllgysysrlesgek
            + vtL+C+          + v y + k ++          r++ +
ht268   41   EKPVTLRQCQP-----PGVDLY-RLEK1SSS-----RYQDQ-- 70

            anlsegrfssissltLtissvekeDsGtYtCvv<-*
                      ++L i   +++ +G Y+C
ht268   71   -----AVLFIPAMKRSLAGRYCSY   90
  
```

FIG.5A

```

      *->GesvtLtCsvgsgfgppgsvtWyfkngk.lgpsllgysysrlesgek
            G++vtL+C+++      + ++ y k+g++ +      y+++
ht268   127   GGDVTLQCQTR---YGFQQFALY-KEGDpAP-----YKNPERWYR-- 162

            anlsegrfssissltLtissvekeDsGtYtCvv<-*
                      +++++i+++v++ sGtY+C
ht268   163   -----ASFPIITVTAHSGTYRCYS   182
  
```

FIG.5B





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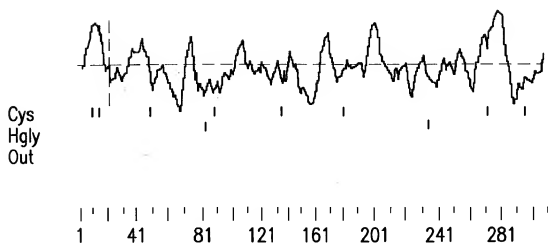
	M	S	P	A	4
GAGTCGACCCACGCGTCCGCTTCCCTGCTTGGCCACATAGCTCAGGACTGGGTTGCAGAACC	ATG	TCT	CCA	GCC	74
S P T F F C I G L C V L Q V I Q T Q S G					24
TCA CCC ACT TTC TTC TGT ATT GGG CTG TGT GTA CTG CAA GTG ATC CAA ACA CAG AGT GGC					134
P L P K P S L Q A Q P S S L V P L G Q S					44
CCA CTC CCC AAG CCT TCC CTC CAG GCT CAG CCC AGT TCC CTG GTA CCC CTG GGT CAG TCA					194
V I L R C Q G P P D V D L Y R L E K L K					64
GTT ATT CTG AGG TGC CAG GGA CCT CCA GAT GTG GAT TTA TAT CGC CTG GAG AAA CTG AAA					254
P E K Y E D Q D F L F I P T M E R S N A					84
CCG GAG AAG TAT GAA GAT CAA GAC TTT CTC TTC ATT CCA ACC ATG GAA AGA AGT AAT GCT					314
G R Y R C S Y Q N G S H W S L P S D Q L					104
GGA CGG TAT CGA TGC TCT TAT CAG AAT GGG AGT CAC TGG TCT CTC CCA AGT GAC CAG CTT					374
E L I A T G V Y A K P S L S A H P S S A					124
GAG CTA ATT GCT ACA GGT GTG TAT GCT AAA CCC TCA CTC TCA GCT CAT CCC AGC TCA GCA					434
V P Q G R D V T L K C Q S P Y S F D E F					144
GTC CCT CAA GGC AGG GAT GTG ACT CTG AAG TGC CAG AGC CCA TAC AGT TTT GAT GAA TTC					494
V L Y K E G D T G P Y K R P E K W Y R A					164
GTT CTA TAC AAA GAA GGG GAT ACT GGG CCT TAT AAG AGA CCT GAG AAA TGG TAC CGG GCC					554
N F P I I T V T A A H S G T Y R C Y S F					184
AAT TTC CCC ATC ATC ACA GTG ACT GCT GCT CAC AGT GGG ACG TAC CGG TGT TAC AGC TTC					614
S S S S P Y L W S A P S D P L V L V V T					204
TCC AGC TCA TCT CCA TAC CTG TGG TCA GCC CCG AGT GAC CCT CTA GTG CTT GTG GTT ACT					674
G L S A T P S Q V P T E E S F P V T E S					224
GGA CTC TCT GCC ACT CCC AGC CAG GTA CCC ACG GAA TCA TTT CCT GTG TAC GAA TCC					734
S R R P S I L P T N K I S T T E K P M N					244
TCC AGG AGA CCT TCC ATC TTA CCC ACA AAC AAA ATA TCT ACA ACT GAA AAG CCT ATG AAT					794
I T A S P E G L S P P I G F A H Q H Y A					264
ATC ACT GCC TCT CCA GAG GGG CTG AGC CCT CCA ATT GGT TTT GCT CAT CAG CAC TAT GCC					854
K G N L V R I C L G A T I I I I L L G L					284
AAG GGG AAT CTG GTC CGG ATA TGC CTT GGT GCC ACG ATT ATA ATA ATT TTG TTG GGG CTT					914
L A E D W H S R K K C L Q H R M R A L Q					304
CTA GCA GAG GAT TGG CAC AGT CGG AAG AAA TGC CTG CAA CAC AGG ATG AGA GCT TTG CAA					974
R P L P P L P L A *					314
AGG CCA CTA CCA CCC CTC CCA CTG GCC TAG					1004
AAATAACTTGGCTTTTCAGCAGAGGGGATTGACCGACATCCATGCACAACCATGGACATACCACTAGAGCCACAGACAT					1083
GGACATACTCAAGAGTGGGAGGTTATATAAAAAAATGAGTGTGAGAAATAAATGCAGAGCCAACAAGGTGAAAAAAA					1162
A					1163

FIG.6



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MSPASPTFFCIGLCVLQVIQTQSGPLPKPSLQAQPSSLVPLGQSVILRCQPPDVLRYL  
EKLKPEKYEDQDFLF IPTMERSNAGRYRCSYQNGSHWSLPDQLEL IATGVYAKPSLSAH  
PSSAVPQGRDVT LKCSQSPYSFDEFVLYKEGDTGPYKRPEKWYRANFPIITVTAHSGTYR  
CYSFSSSSPYLWSAPSDPLVL VVTGLSATPSQVPTEESFPVTESSRRPSILPTNKISTTE  
KPMNITASPEGLSPPIGFAHQHYAKGNLVRICLGATIIIIILLGLLAEDWHSRKKCLQHRM  
RALQRPLPPLPLA

FIG.7



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      10      20      30      40      50      60      70
inputs  ATGACGCCGCCCTCACAGCCCTGCTGTGCCTTGGGCTGAGCTGGGCCCCAGGACCCGCGTGCAGGCAG
      : : : : : : : : : : : : : : : : : : : : : :
      ATGTCCCAGCC--TCAC--CC---ACTTTCCT--CTGTAT-----
      10      20      30

      80      90      100      110      120      130      140
inputs  GGCCTTCCCCAACCCACCCCTCTGGGCTGAGCCAGGCTCTGTGATCAGCTGGGGAGCCCCGTGACCAT
      : : : : : : : : : : : : : : : : : : : : : :
      -----TGGGCTG-----TGTTACTGC-----
                        40

      150      160      170      180      190      200      210
inputs  CTGGTGTCAAGGGAGCCTGGAGGCCAGGAGTACCGACTGGATAAAGAGGGAAGCCAGAGCCCTGGAC
      : : : : : : : : : : : : : : : : : : : : : :
      -----AAGTGATCC-----AAACACAGAG---TGG--
                        50                        60                        70

      220      230      240      250      260      270      280
inputs  AGAATAACCCACTGGAACCCCAAGAACAGGCCAGATTCTCCATCCATCCATGACAGAGCACCATTGCGG
      : : : : : : : : : : : : : : : : : : : : : :
      -----CCCACT---CCC---CAAG-----CCTTCCC--TCCAGG-----
                        80                        90

      290      300      310      320      330      340      350
inputs  GGAGATACCGCTGCCACTATTACAGCTCTGCAGGCTGGTCAGAGCCAGCGACCCCTGGAGCTGGTGAT
      : : : : : : : : : : : : : : : : : : : : : :
      -----CTCAGCC-----CAGTCCCTG--GTACCCCTGGGTGAG-----
      100                        110                        120

      360      370      380      390      400      410      420
inputs  GACAGGATTCTACAACAACCCACCCCTCTCAGCCCTGCCAGCCCTGTGGTGGCTCAGGGGGGAATATG
      : : : : : : : : : : : : : : : : : : : : : :
      -TCAG--TTATTC-----TGAGGTG-C--CAGGGA-----
      130                        140                        150

      430      440      450      460      470      480
inputs  ACCCTCC--GATGTGGCTACAGAAGGGATATCACCATTITGTTCTGATGAAGGAAGGAGAACCAGCTC
      : : : : : : : : : : : : : : : : : : : : : :
      -CCTCCAGATGTGG-----ATTTATATCGCCTGGAGAAACTGAAA-----
      160      170      180      190

      490      500      510      520      530      540      550
inputs  CCCCAGACCTGGACTCACAGAGCTCCACAGTGGGGGTTCCAGGCCCTGTTCCCTGTGGGCCCCGTGA
      : : : : : : : : : : : : : : : : : : : : : :
      --CCGGA---GA-----AGTATGAAGATCAAGAC---TTTCTCTT-----CAIT-
                        200      210      220
```

FIG. 8A

FIG. 8B



```

1120      1130      1140      1150      1160      1170      1180
inputs  GCTCATAAGTACCAAGGCTGAATTCCCCATGAGTCCTGTGACCTCAGCCACGCGGGACCTACAGGTGCT
      : .....: .....: .....: .....: .....: .....: .....:
      G--AAATGGTACCGGGCCAATTTCCCATCATCACAGTGACTGCTGCTACAGTGGGACGTACCGGTGTT
      480      490      500      510      520      530      540

1190      1200      1210      1220      1230      1240      1250
inputs  ACGGCTCATACAGCTCCAACCCCCACCTGCTGTCTTTCCCAAGTGAGCCCTGGAACTCATGGTCTCAGG
      : .....: .....: .....: .....: .....: .....: .....:
      ACAGCTTCTCCAGCTCATCTCCATACCTGTGGTCAGCCCGAGTGACCCTCTAGTGTCTGTGGTTACTGG
      550      560      570      580      590      600      610

1260      1270      1280      1290      1300      1310      1320
inputs  ACACTCTGGAGGCTCCAGCCTCCCACCCACAGGGCCGCCCTCCACACCTGGTCTGGGAAGATACCTGGAG
      : .....: .....: .....: .....: .....: .....: .....:
      ACTCTCTG-----CCA--CTCCAGCC--AGGT--ACCCAC-----GGA--AGAATCATTTCTG---
      620      630      640      650      660

1330      1340      1350      1360      1370      1380      1390
inputs  GTTTGATTGGGGCTCTCGGTGGCCTTCGTCTGCTGCTTCTCTCTCTCTCTCTCTCTCGACGTC
      : .....: .....: .....: .....: .....: .....: .....:
      ----TGA-----CAGAACTCT--CCAGGAGACCTTCCA-----TCTTAC----CCACAAACAA
      670      680      690      700

1400      1410      1420      1430      1440      1450      1460
inputs  AGCGTCACAGCAAAACAGGACATCTGACCAGAGAAAGACTGATTTCAGCGTCTGCAAGGGGCTCGGGA
      : .....: .....: .....: .....: .....: .....: .....:
      A---TATCTACAA---CTGAA---AAGCCTATGAATATC--ACTGCCT-C-TCCAG-AGGGGCTG---
      710      720      730      740      750

1470      1480      1490      1500      1510      1520      1530
inputs  GACAGAGCCCAAGGACAGGGGCCTGCTGAGGAGGTCCAGCCAGCTGTGACGTCCAGGAAGAAAACCTC
      : .....: .....: .....: .....: .....: .....: .....:
      ----AGCCCT-----CC-----AATTGGTTTGTCTCATCAGCA-----C
      760      770      780

1540      1550      1560      1570      1580      1590      1600
inputs  TATGTCGCGTGAAGGACACACAGTCTGAGGACAGGGTGGAGCTGGACAGTCAGAGCCACACGATGAAG
      : .....: .....: .....: .....: .....: .....: .....:
      TATGC-----CAAGGGGAATCTGGTC-----CGGATATG
      790      800      810

1610      1620      1630      1640      1650      1660      1670
inputs  ACCCCACAGGACGTGACGTATGCCCGGTGAAACACTCAGTCTTAGGAGAGAAATGGCTCTCTCCCTC
      : .....: .....: .....: .....: .....: .....: .....:
      --CCTTGG-----TGCCACGAT-----TATAATAATTTTGT-----
      820      830      840

1680      1690      1700      1710      1720      1730      1740
inputs  CTCACTGTCTGGGGAATTCCTGGACACAAAGGACAGACAGGTGGAAGAGGACAGGCAGATGGACACTGAG
      : .....: .....: .....: .....: .....: .....: .....:
      -----TGGGGCTT--CTAG--CAGAGGATTGGC-----ACAGTCGGAAGAA--AT
      850      860      870      880
```

FIG.8C



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```
      1750      1760      1770      1780      1790      1800      1810
inputs  GCTGCTGCATCTGAAGCCTCCCAGGATGTGACCTACGCCAGCTGCACAGCTTGACCTTAGACGGAAGG
      :: :::::.:: :::::.:: :::::.:: :::::.::
      GC--CTGCAACA-----CAGGATGAGA-----GCTTTGC-----AAAGG
                890                900                910
```

```
      1820      1830      1840      1850      1860      1870      1880
inputs  CAACTGAGCCTCCTCCATCCCAGGAAGGGGAACCTCCAGCTGAGCCAGCATCTACGCCACTCTGGCCAT
      : :::.:: :::::.:: :::::.:: :::::.::
      CCACTA-----CCACC-----CCTCC-----CACTGGCC--
                920                930
```

```
      1890
inputs  CCAC
```

FIG. 8D



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```

      10      20      30      40      50      60
inputs MSPASPTFFCIGLCVLQVIQTQSGPLPKPSLQAQPSLVPLGQSVILRCQGPDPVDLYRLEKL-KPEKYE
      ..
      MTPALTALLCLGLSLGPRTRVQAGFPFKPTLWAEPGSVISWGSPTTIWCOGSLEAQEYRLDKEGSPPELD
      10      20      30      40      50      60      70
      70      80      90      100      110      120      130
inputs DQDFL-----F-IPTMERSNAGRYRCSYQNGSHWSLPSDQLELIATGVYAKPSLSAHPSSAVPQGRDV
      ..
      RNNPLEPKNKARFISPMTEHHAGRYRCHYYSSAGNSEPSDPLELVMGTGFYINKPTLSALPSPVVASGGM
      80      90      100      110      120      130      140

inputs TLKC--QSPY-----
      ..
      TLRCGSKQGYHHFVLMKEGEHQLPRTLDSQQHLHSGGFQALFPVGPVNPNSHRWRFTCYYYMMNTPQVWSHP
      150      160      170      180      190      200      210

inputs -----140      150-----
      ..
      SDPLEILPSGVSRRKPSLLTLQGPVLAPGQSLTLQCSDVGYYDRFVLYKEGERDFLQRGQQPAGLSQAN
      220      230      240      250      260      270      280

inputs -----160-----
      ..
      FTLGPVSPSHGGQYRCYGAHNLSSEWSAPSDDLILMAGQIYDTVSLSAQPGPTVASGENVTLLCQSMWQ
      290      300      310      320      330      340      350

inputs -----170      180      190      200-----
      ..
      YRANFPIITVTAHSGTYRCYSFSSSPYLWSAPSDDLVLVVTG
      170      180      190      200
      FDTLLTKEGAHPPLRLRSMYGAHKYQAEFPMSPVTSAHAGTYRCYGSYSSNPHELLFPSEPLELMVSG
      360      370      380      390      400      410      420

inputs -----210      220-----
      ..
      LSATPSQVPTES-----FPV-----
      210      220
      HSGGSSLPPTGPPSTPGLGRYLEVLIGVSVAFLVLLFLLFLLRQRQHSKHTSDQRKTDQFQAGAAE
      430      440      450      460      470      480      490

input TESS-----RRPS-----230      240      250      260      270
      ..
      ILPTNKISTTEKPMNI-TASPEGLSP-PIGFAH--QHYAKGNLVR--I
      230      240      250      260      270
      TEPKDRGLLRRSSPAADVQENLYAAVKDTQSEDRLVDSQSPHDEDPQAVTYAPVKHSPPREMASPPS
      500      510      520      530      540      550      560

inputs -----280      290      300      310-----
      ..
      CLGATIIIIILLGLLAEDWH-----SRKKCLQHRMRALQRPL-----LPL
      280      290      300      310
      SLSGEFLDTKDRQVEEDROMDEAAASEASQDVTYAQLHSLTLRRKATEPPSQEGEPPEAPESYATLAI
      570      580      590      600      610      620      630

inputs A
H
```

FIG. 9



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```

*->GesvtLtCsvgfgppgvsvtWyfkngk.lgpsllgysysrlesgek
G+sv L+C+      ++v y + k ++      +++e +
mT268  42  GQSVILRCQGP-----PDVDLY-RLEK1KP-----EKYEDQ-- 71

anlsegrfsissltLtissvekeDsGtYtCvv<-*
L i + e++++G Y+C
mT268  72  -----DFLFIPTMERSNAGRYRCSY 91
  
```

FIG.10A

```

*->GesvtLtCsvgfgppgvsvtWyfkngk.lgpsllgysysrlesgek
G +vtL C++      ++ y k+g++ +      Y+r+e +
mT268  128  GRDVTLCQSP---YSFDEFVLY-KEGdtGP-----YKRPEKW-Y 162

anlsegrfsissltLtissvekeDsGtYtCvv<-*
+      ++i++v++ sGtY+C
mT268  163  RA-----NFPIITVTAHSGTYRCYS 183
  
```

FIG.10B





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	10	20	30	40	50	60	
inputs	MSPSPTALFCLGLCLGRV-PAQSGPLPKPSLQALPSSLVPLEKPVTLRCQGPVGVLYRLEKLSSSRYQD						
	MSPASPTFFCTGLCVLQVIQTQSGPLPKPSLQALPSSLVPLGQSVILRCQGPVGVLYRLEKLSPEKYED						
	10	20	30	40	50	60	70
inputs	QAVLFIPAMKRSAGRYRCSYQNSLWSLPSDQLELVATGVFAKPSLSAQPGPAVSSGGDVTLCQTRYG						
	QDFLFIPTMERSNAGRYRCSYQNSHWSLPSDQLELVATGVYAKPSLSAHPSSAVPGROVTLKCQSPYS						
	80	90	100	110	120	130	140
inputs	FDQFALYKEGDPAKYKNPERWYRASFPITVTAAHSGTYRCYSFSSRDPLYWSAPSDPLELVVTGTSVTP						
	FDEFVLYKEGDTGPKRPEKMYRANFPITVTAAHSGTYRCYSFSSSPPLYWSAPSDPLVLVVTGLSATP						
	150	160	170	180	190	200	210
inputs	SRLPTEPPSSVAEFSEATAELTVSFNKKVFTTETSRSITTSPEKESDPAGPARQYTYKGNLVRLCLGAVI						
	SQVPTEESFPVTSSRRPSILP---TNKISTTEKPMNITASPEGLSPPIGFAHQHYAKGNLVRLCLGATI						
	220	230	240	250	260	270	
	280	290	300	310	320	330	
inputs	LIILAGFLAEDWHSRKRRLRHRGRAVQRPLPLPPLPQTRKSHGGQDGRQDVHSRGLCS						
	IIILLGLLAEDWHSRKKCLQHRMRALQRLPLP-LA-----						
	280	290	300	310			

FIG.11

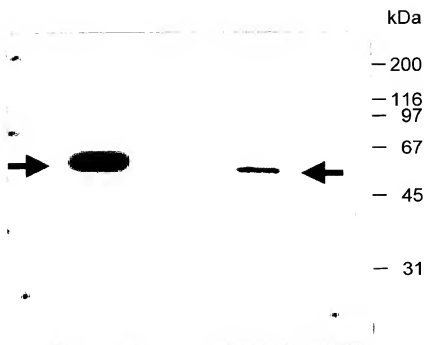
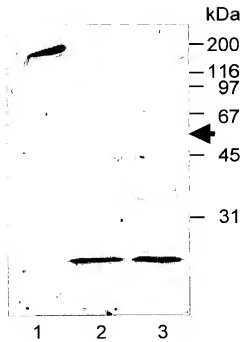
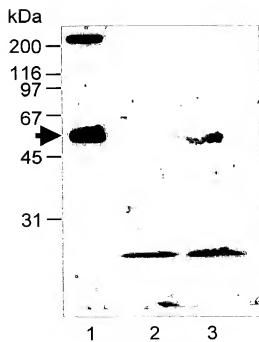


FIG.12

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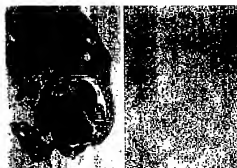


FIG.14A

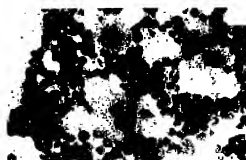


FIG.14B

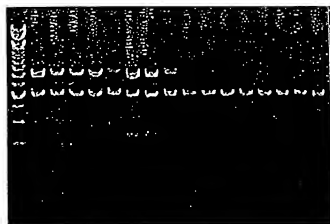


FIG.14C

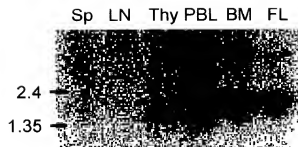


FIG.14D

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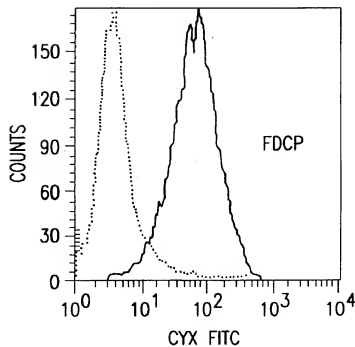


FIG.15A

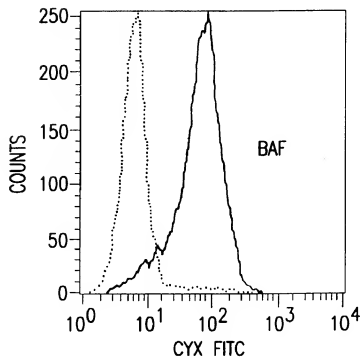


FIG.15B



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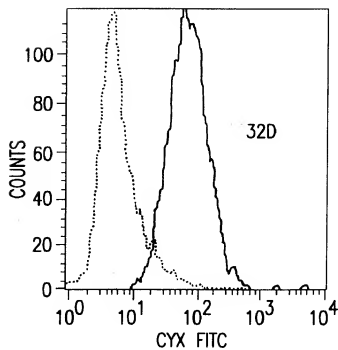


FIG.15C

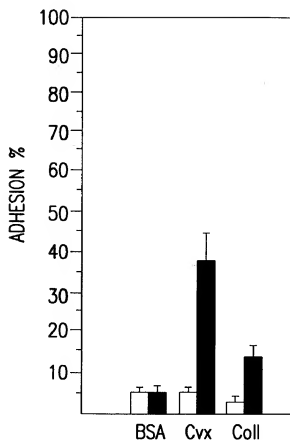


FIG. 16A

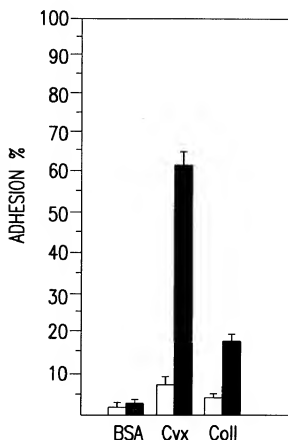


FIG. 16B



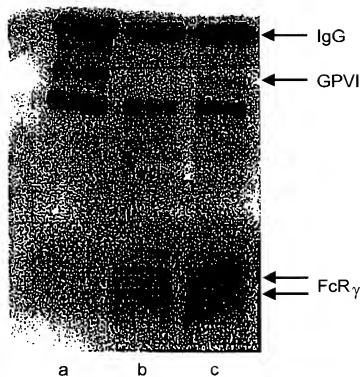


FIG.17

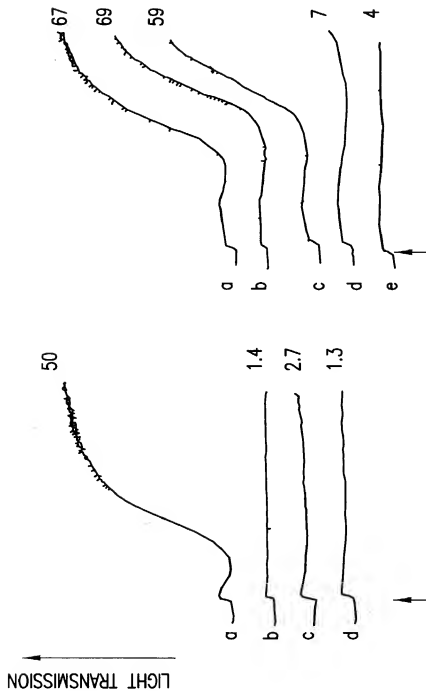


FIG. 18B

FIG. 18A



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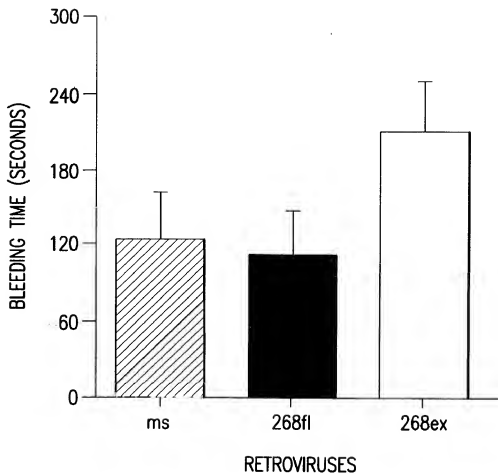


FIG. 19

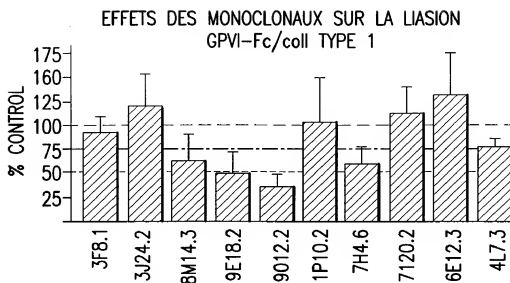


FIG.20

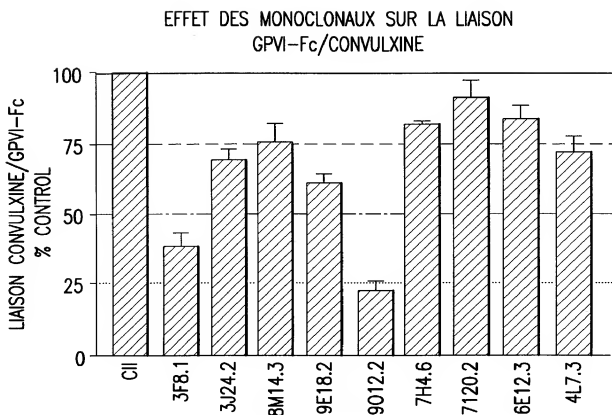


FIG.21



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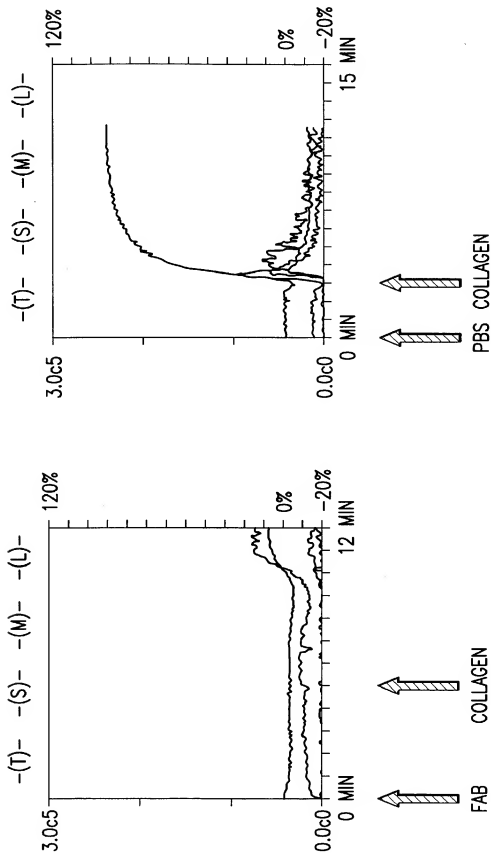


FIG.22

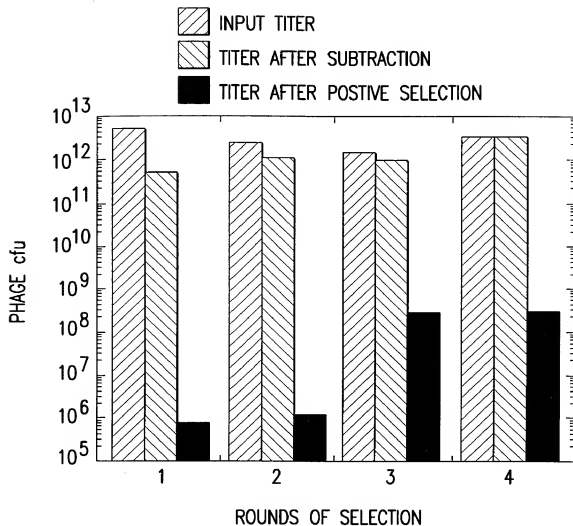


FIG.23



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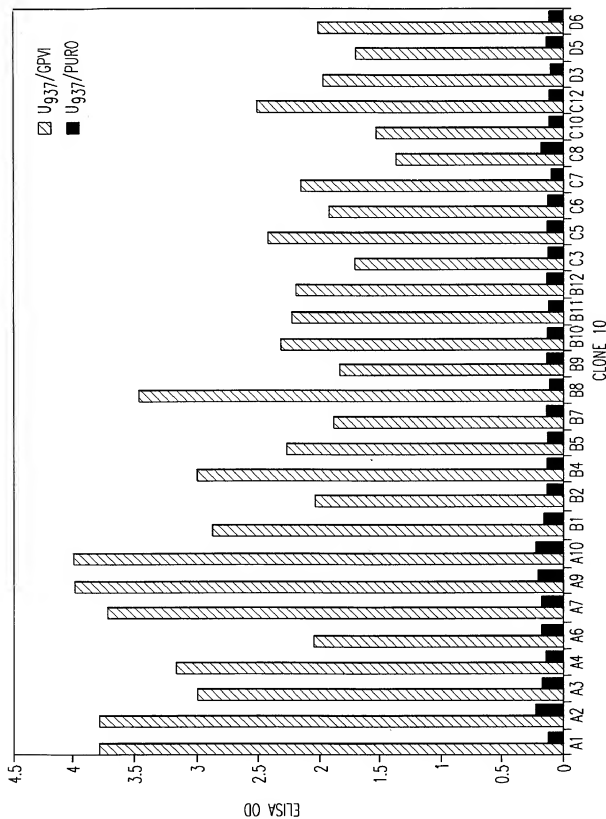


FIG.24A



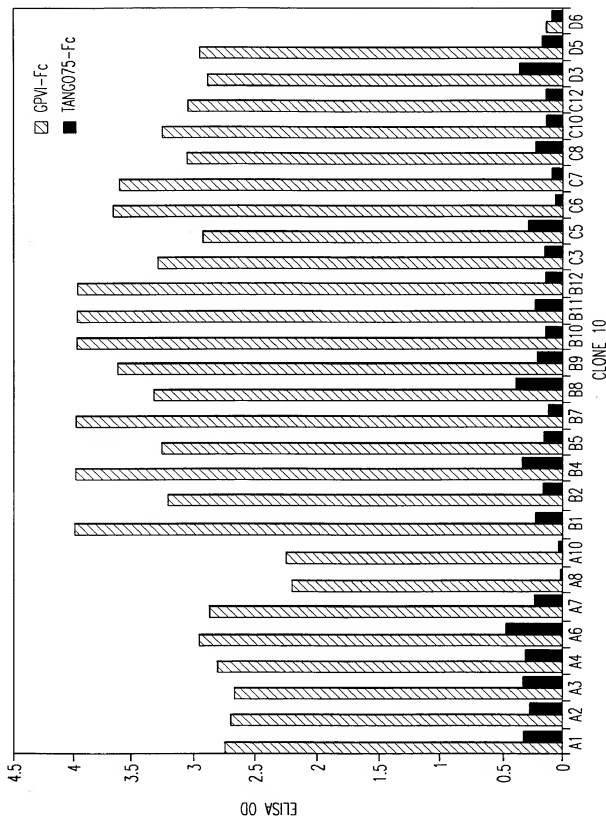


FIG. 24B

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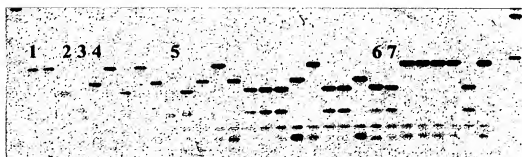


FIG.25



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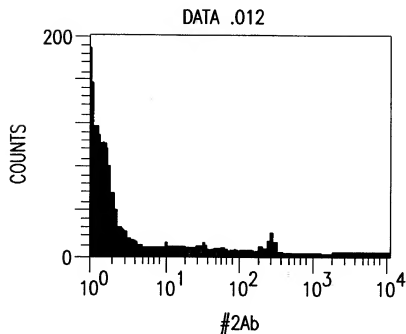


FIG.26A

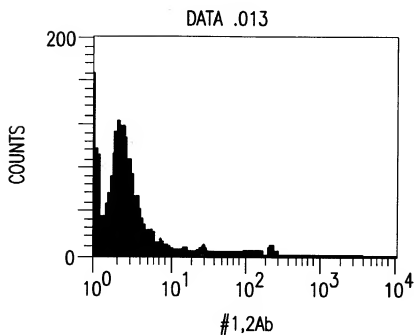


FIG.26B



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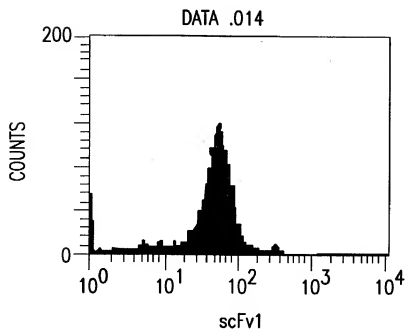


FIG.26C

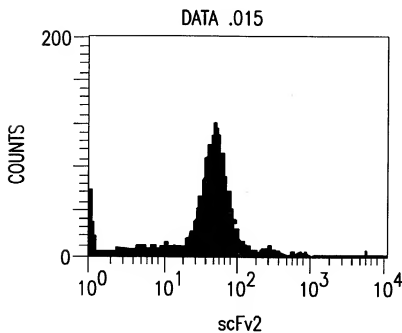


FIG.26D

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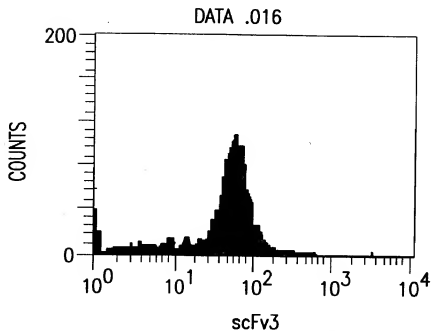


FIG.26E

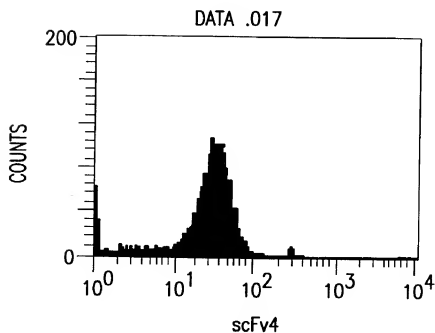


FIG.26F



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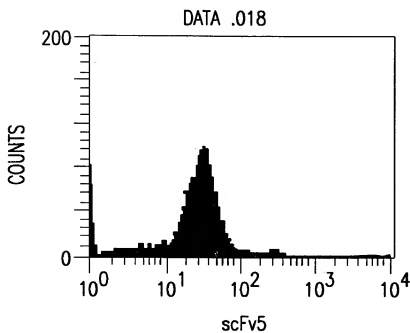


FIG.26G

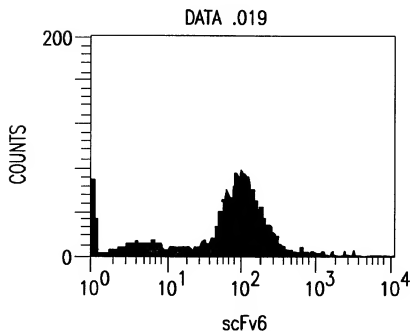
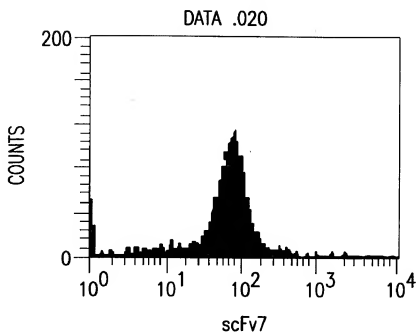


FIG.26H



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scFv1:A4  
scFv2:B4  
scFv3:A9  
scFv4:C4  
scFv5:C9  
scFv6:C10  
scFv7:A10

FIG.26I

20/T0/80



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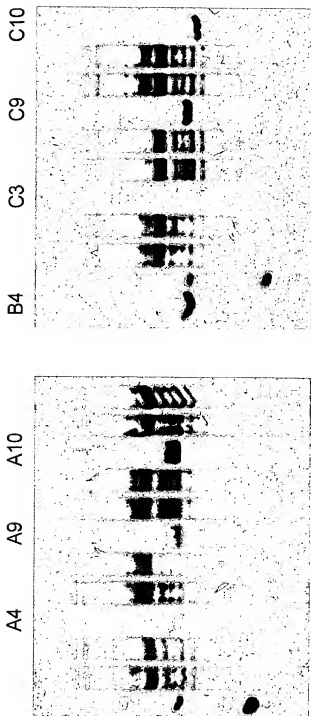


FIG.27

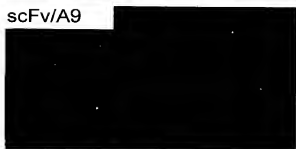




control scFv



scFv/A9



scFv/C3

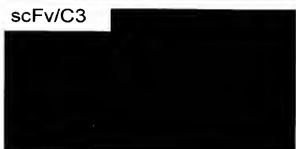


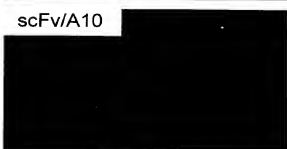
FIG.28A



scFv/A4



scFv/A10



scFv/C9



FIG.28B

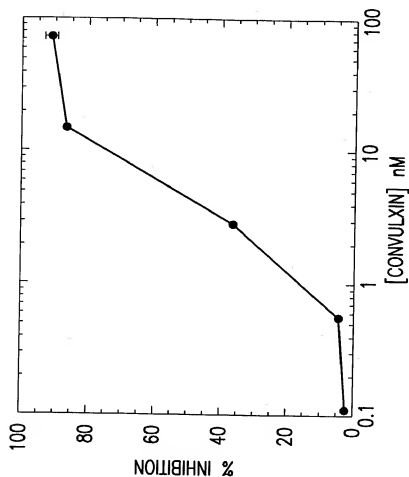


FIG. 29B

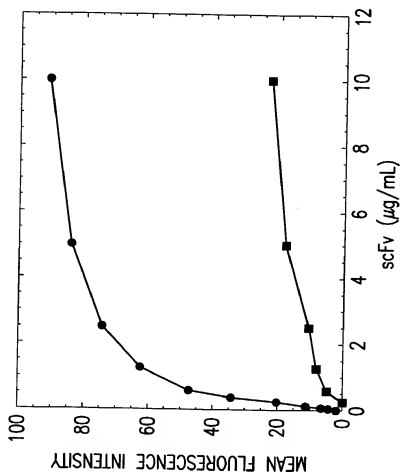


FIG. 29A

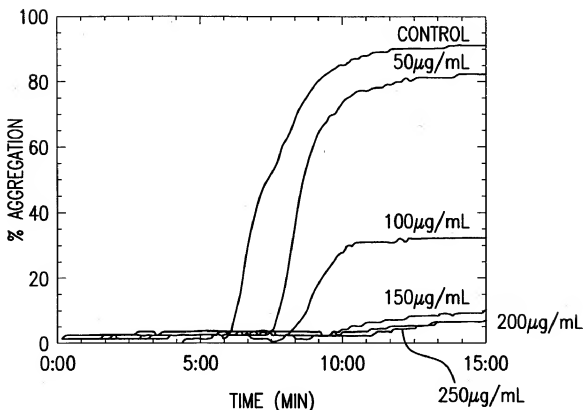


FIG.30A

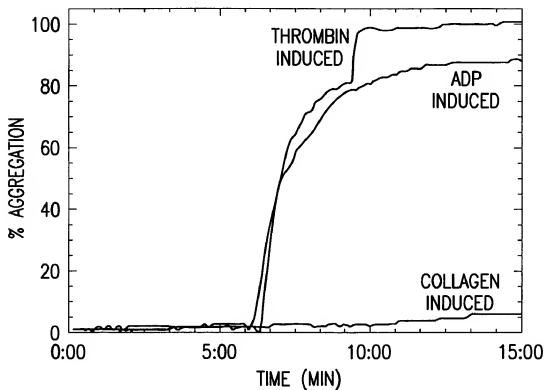


FIG.30B